ABSTRACT

TRIAZOLOPYRIMIDINE DERIVATIVES AS GLYCOGEN SYNTHASE KINASE 3 INHIBITORS

This invention concerns a compound of formula

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$$\begin{array}{c|c}
R^3 & R^1 & X \\
 & X$$

a *N*-oxide, a pharmaceutically acceptable addition salt, a quaternary amine and a stereochemically isomeric form thereof, wherein ring A represents phenyl, pyridyl, pyrimidinyl, pyridazinyl or pyrazinyl; R¹ represents hydrogen; aryl; formyl; C₁₋₆alkylcarbonyl; C₁₋₆alkylcycarbonyl; C₁₋₆alkyls substituted with formyl, C₁₋₆alkylcarbonyl, C₁₋₆alkyloxycarbonyl, C₁₋₆alkylcarbonyloxy; or C₁₋₆alkylcycarbonyl optionally substituted with C₁₋₆alkyloxycarbonyl; X represents a direct bond; -(CH₂)_{n3}-or -(CH₂)_{n4}-X_{1a}-X_{1b}-; R² represents C₃₋₇cycloalkyl; phenyl; a 4, 5, 6- or 7-membered monocyclic heterocycle containing at least one heteroatom selected from O, S or N; benzoxazolyl or a radical of formula

wherein said R² substituent may optionally be substituted; R³ represents halo; hydroxy; optionally substituted C₁₋₆alkyl; C₂₋₆alkenyl or C₂₋₆alkynyl, each optionally substituted; optionally substituted polyhaloC₁₋₆alkyl; optionally substituted C₁₋₆alkyloxy; optionally substituted polyhaloC₁₋₆alkyloxy; C₁₋₆alkylthio; polyhaloC₁₋₆alkylthio; C₁₋₆alkyl-oxycarbonyl; C₁₋₆alkylcarbonyloxy; C₁₋₆alkylcarbonyl; polyhaloC₁₋₆alkylcarbonyl; cyano; carboxyl; aryloxy; arylthio; arylcarbonyl; NR^{6b}R^{7b}; C(=O)-NR^{6b}R^{7b}; -NR⁵-C(=O)-R⁵; -S(=O)_{n1}-R^{8a}; -NR⁵-S(=O)_{n1}-R^{8a}; -S-CN; -NR⁵-CN; R⁴ represents hydrogen; halo; hydroxy; optionally substituted C₁₋₄alkyl; C₂₋₄alkenyl or C₂₋₄alkynyl, each optionally substituted; polyhaloC₁₋₃alkyl; optionally substituted C₁₋₄alkyloxy; polyhalo-C₁₋₃alkyloxy; C₁₋₄alkylthio; polyhaloC₁₋₃alkylthio; C₁₋₄alkyloxycarbonyl; C₁₋₄alkylcarbonyloxy; C₁₋₄alkylcarbonyl; polyhaloC₁₋₄alkylcarbonyl; nitro; cyano; carboxyl; NR¹⁰R¹¹; C(=O)NR¹⁰R¹¹; -NR⁵-C(=O)-NR¹⁰R¹¹; -NR⁵-C(=O)-R⁵; -S(=O)_{n1}-R¹²; -NR⁵-S(=O)_{n1}-R¹²; -S-CN; -NR⁵-CN; their use, pharmaceutical compositions comprising them and processes for their preparation.